

REMARKS

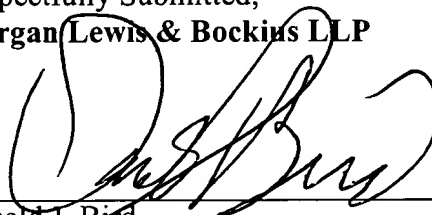
The above amendment to the claim 1 is for the sole purpose of conforming the claims to the elected invention. Claim 11 has been cancelled as being in a "use" format inappropriate to U.S. practice. Pharmaceutical composition claim 10 has been amended to make it dependent on any one of claims 1-8. Method of treatment claim 12 has been amended and new claim 13 has been added to separate the recitations in original claim 12, of compounds of claim 1 and the specifically named compound, into separate claims. Claim 12 has also been amended to be dependent on any one of claims 1-8, and to be in a more customary format for U.S. practice. The amendments are made without waiver or prejudice to Applicant's right to prosecute the non-elected subject matter in one or more divisional applications. Entry of these amendments and a favorable action on these claims is respectfully requested.

The Examiner's attention is also called to the Information Disclosure Statement being filed herewith. It is respectfully requested that the reference accompanying this Information Disclosure Statement be considered when this application is acted on, and that such consideration be acknowledged by initialing and returning a copy of the accompanying Form PTO-1449. The Examiner is also reminded to take into consideration (and acknowledge such consideration) of the documents cited in the International Search Report and PTO-1449 filed with this application on September 20, 2001, and the related applications and documents cited in and filed with the Information Disclosure Statement and PTO-1449, filed and dated January 3, 2003.

Conclusion

Entry of the above amendments and a favorable action on the merits are respectfully requested, together with consideration of the cited documents of record in this application.

Respectfully Submitted,
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VERSION WITH MARKINGS TO SHOW CHANGES

1-WA/1931221.1



wherein A is halo, hydroxy, C₁₋₆alkoxy, C₁₋₆alkylS(O)_n- (wherein n is 0-2), cyano, amino, *N*-C₁₋₆alkylamino, *N,N*-(C₁₋₆alkyl)₂amino, carboxy, C₁₋₆alkoxycarbonyl, carbamoyl, *N*-C₁₋₆alkylcarbamoyl or *N,N*-(C₁₋₆alkyl)₂carbamoyl, p is 1 - 6, and B is a bond, oxy, imino, *N*-(C₁₋₆alkyl)imino or -C(O)NH-, with the proviso that p is 2 or more unless B is a bond or -C(O)NH-,

or R¹ is of the Formula (IB):



wherein D is aryl, heteroaryl or heterocyclyl and E is a bond, C₁₋₆alkylene, C₁₋₆alkyleneoxy, oxy, imino, *N*-(C₁₋₆alkyl)imino, C₁₋₆alkyleneimino, *N*-(C₁₋₆alkyl)-C₁₋₆alkyleneimino, C₁₋₆alkyleneoxy-C₁₋₆alkylene, C₁₋₆alkyleneimino-C₁₋₆alkylene, *N*-(C₁₋₆alkyl)-C₁₋₆alkyleneimino-C₁₋₆alkylene, -C(O)NH-, -SO₂NH-, -NH₂SO₂- or C₂₋₆alkanoylimino,

and any aryl, heteroaryl or heterocyclyl group in a R¹ group may be optionally substituted with one or more groups selected from hydroxy, halo, C₁₋₆alkyl, C₁₋₆alkoxy, carboxy, C₁₋₆alkoxycarbonyl, carbamoyl, *N*-C₁₋₆alkylcarbamoyl, *N,N*-(C₁₋₆alkyl)₂carbamoyl, C₂₋₆alkanoyl, amino, *N*-C₁₋₆alkylamino and *N,N*-(C₁₋₆alkyl)₂amino,

and any heterocyclyl group in a R¹ group may be optionally substituted with one or two oxo or thioxo substituents,

and any of the R¹ groups defined hereinbefore which comprises a CH₂ group which is attached to 2 carbon atoms or a CH₃ group which is attached to a carbon atom may optionally bear on each said CH₂ or CH₃ group a substituent selected from hydroxy, amino, C₁₋₆alkoxy, *N*-C₁₋₆alkylamino, *N,N*-(C₁₋₆alkyl)₂amino and heterocyclyl;

R² is hydrogen, halo, C₁₋₆alkyl, C₂₋₆alkenyl or C₂₋₆alkynyl;

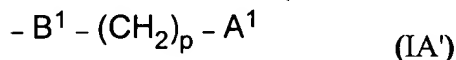
R³ is hydrogen, halo, C₁₋₆alkyl, C₂₋₆alkenyl or C₂₋₆alkynyl;

R⁴ is hydrogen, hydroxy, C₁₋₆alkyl, C₁₋₆alkoxy, amino, *N*-C₁₋₆alkylamino, *N,N*-(C₁₋₆alkyl)₂amino, hydroxyC₂₋₆alkoxy, C₁₋₆alkoxyC₂₋₆alkoxy, aminoC₂₋₆alkoxy, *N*-C₁₋₆alkylaminoC₂₋₆alkoxy, *N,N*-(C₁₋₆alkyl)₂aminoC₂₋₆alkoxy or C₃₋₇cycloalkyl,

or R⁴ is of the Formula (IC):



wherein J is aryl, heteroaryl or heterocyclyl and K is a bond, oxy, imino, *N*-(C₁₋₆alkyl)imino, oxyC₁₋₆alkylene, iminoC₁₋₆alkylene, *N*-(C₁₋₆alkyl)iminoC₁₋₆alkylene, -NHC(O)-, -SO₂NH-, -NHCO₂- or -NHC(O)-C₁₋₆alkylene-, and any aryl, heteroaryl or heterocyclyl group in a R⁴ group may be optionally substituted by one or more groups selected from hydroxy, halo, trifluoromethyl, cyano, mercapto, nitro, amino, carboxy, carbamoyl, formyl, sulphamoyl, C₁₋₆alkyl, C₂₋₆alkenyl, C₂₋₆alkynyl, C₁₋₆alkoxy, -O-(C₁₋₃alkyl)-O-, C₁₋₆alkylS(O)_n- (wherein n is 0-2), *N*-C₁₋₆alkylamino, *N,N*-(C₁₋₆alkyl)₂amino, C₁₋₆alkoxycarbonyl, *N*-C₁₋₆alkylcarbamoyl, *N,N*-(C₁₋₆alkyl)₂carbamoyl, C₂₋₆alkanoyl, C₁₋₆alkanoyloxy, C₁₋₆alkanoylamino, *N*-C₁₋₆alkylsulphamoyl, *N,N*-(C₁₋₆alkyl)₂sulphamoyl, C₁₋₆alkylsulphonylamino and C₁₋₆alkylsulphonyl-*N*-(C₁₋₆alkyl)amino, or any aryl, heteroaryl or heterocyclyl group in a R⁴ group may be optionally substituted with one or more groups of the Formula (IA'):



wherein A¹ is halo, hydroxy, C₁₋₆alkoxy, cyano, amino, *N*-C₁₋₆alkylamino, *N,N*-(C₁₋₆alkyl)₂amino, carboxy, C₁₋₆alkoxycarbonyl, carbamoyl, *N*-C₁₋₆alkylcarbamoyl or *N,N*-(C₁₋₆alkyl)₂carbamoyl, p is 1 - 6, and B¹ is a bond, oxy, imino, *N*-(C₁₋₆alkyl)imino or -NHC(O)-, with the proviso that p is 2 or more unless B¹ is a bond or -NHC(O)-, or any aryl, heteroaryl or heterocyclyl group in a R⁴ group may be optionally substituted with one or more groups of the Formula (IB'):



wherein D¹ is aryl, heteroaryl or heterocyclyl and E¹ is a bond, C₁₋₆alkylene, oxyC₁₋₆alkylene, oxy, imino, *N*-(C₁₋₆alkyl)imino, iminoC₁₋₆alkylene, *N*-(C₁₋₆alkyl)iminoC₁₋₆alkylene, C₁₋₆alkylene-oxyC₁₋₆alkylene, C₁₋₆alkylene-iminoC₁₋₆alkylene, C₁₋₆alkylene-*N*-(C₁₋₆alkyl)iminoC₁₋₆alkylene, -NHC(O)-, -NHCO₂-, -SO₂NH- or -NHC(O)-C₁₋₆alkylene-, and any aryl, heteroaryl or heterocyclyl group in a substituent on R⁴ may be optionally substituted with one or more groups selected from hydroxy, halo, C₁₋₆alkyl, C₁₋₆alkoxy,

carboxy, C₁₋₆alkoxycarbonyl, carbamoyl, *N*-C₁₋₆alkylcarbamoyl, *N*-(C₁₋₆alkyl)₂carbamoyl, C₂₋₆alkanoyl, amino, *N*-C₁₋₆alkylamino and *N,N*-(C₁₋₆alkyl)₂amino, and any C₃₋₇cycloalkyl or heterocyclyl group in a R⁴ group may be optionally substituted with one or two oxo or thioxo substituents, and any of the R⁴ groups defined hereinbefore which comprises a CH₂ group which is attached to 2 carbon atoms or a CH₃ group which is attached to a carbon atom may optionally bear on each said CH₂ or CH₃ group a substituent selected from hydroxy, amino, C₁₋₆alkoxy, *N*-C₁₋₆alkylamino, *N,N*-(C₁₋₆alkyl)₂amino and heterocyclyl; R⁵ is hydrogen, halo, trifluoromethyl, cyano, nitro, amino, hydroxy, C₁₋₆alkyl, C₂₋₆alkenyl, C₂₋₆alkynyl, C₁₋₆alkoxy, *N*-C₁₋₆alkylamino or *N,N*-(C₁₋₆alkyl)₂amino; q is 0, 1, 2, 3 or 4; or a pharmaceutically acceptable salt or an *in vivo* cleavable ester thereof; with the proviso that 7-amino-4-(3-acetamidoanilino)pyrido[4,3-*d*]pyrimidine is excluded.

10. (Amended) A pharmaceutical composition which comprises a bicyclic compound of the Formula (I), or a pharmaceutically acceptable salt or *in vivo* cleavable ester thereof, according to **[claim 1] any one of claims 1-8** in association with a pharmaceutically acceptable diluent or carrier.

12. (Amended) A method of **[of]** treating **a disease** **[diseases]** or medical **condition** **[conditions]** mediated by cytokines which comprises administering to a warm-blooded animal **in need thereof** an effective amount of a bicyclic compound of the Formula (I), or a pharmaceutically acceptable salt or an *in vivo* cleavable ester thereof, according to **any one of claims 1-8. [claim 1 or of the compound 7-amino-4-(3-acetamidoanilino)pyrido[4,3-*d*]pyrimidine.]**